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7	5	fluoresc\$4 same polari\$7 same estrogen same receptor	USPAT; US-PGPUB; EPO; JPO; DERWENT	2002/08/14 10:59

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L2 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2002:190219 CAPLUS

TITLE: High throughput fluorescence polarization-based screening assays for the identification of novel nuclear receptor ligands

AUTHOR(S): Eliason, Hildegard C.; Shekhani, Mohammed Saleh; Ervin, Kerry M.; Halbleib, Cale M.; Millis, Sherri Z.;

CORPORATE SOURCE: Mei, Baigen; Lowery, Robert G.; Burke, Thomas J. PanVera Corp., Madison, WI, 53719, USA

SOURCE: Abstracts of Papers, 223rd ACS National Meeting, Orlando, FL, United States, April 7-11, 2002 (2002), MEDI-100. American Chemical Society: Washington, D. C.

CODEN: 69CKQP

DOCUMENT TYPE: Conference; Meeting Abstract

LANGUAGE: English

AB **Steroid hormone receptors** (SHRs) are ligand-induced transcription factors that mediate the transactivation of genes responsible for cellular differentiation, reprodn., and metab. PanVera has developed a panel of **fluorescence polarization** (FP)-based high throughput screening assays for the rapid identification of novel SHR ligands for androgen, progesterone, glucocorticoid, and estrogen (alpha and beta) **receptors**. These homogeneous assays utilize recombinant human **receptor** proteins and fluorophore-**steroid** conjugates specific for these **receptors**. The synthetic **fluorescent** ligands bind with affinities similar to that of their resp. native ligands - generally in the low nanomolar range.

In FP assays, the **polarization** of the fluorophore is proportional to the fraction complexed with **receptor**. One can deduce the binding affinity of a test compd. by measuring its ability to

displace a **fluorescent** ligand from the **receptor's** hormone binding pocket. Such screening assays provide a simple and rapid method for detecting novel SHR ligands for this important class of drug targets.

L2 ANSWER 2 OF 8 MEDLINE DUPLICATE 1  
ACCESSION NUMBER: 2000227100 MEDLINE  
DOCUMENT NUMBER: 20227100 PubMed ID: 10766033  
TITLE: Modulation of LH/hCG receptors and physical state of ovarian membranes in rat pseudopregnancy.  
AUTHOR: Jezova M; Scsukova S; Vranova J; Kolena J  
CORPORATE SOURCE: Institute of Experimental Endocrinology, Slovak Academy of Sciences, Bratislava.. ueenjez@savba.savba.sk  
SOURCE: GENERAL PHYSIOLOGY AND BIOPHYSICS, (1999 Dec) 18 (4) 347-56.  
Journal code: 8400604. ISSN: 0231-5882.  
PUB. COUNTRY: Slovakia  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 200005  
ENTRY DATE: Entered STN: 20000606  
Last Updated on STN: 20000606  
Entered Medline: 20000523

AB . . . as well as regression of corpora lutea. The effects of cyclooxygenase inhibitors (indomethacin and acetylsalicylic acid (ASA)) and of selected **steroids** (estradiol, testosterone and dihydrotestosterone) on the functional state of luteinized ovaries were studied. The compounds were administered to the animals. . . .  
injection.  
ASA and indomethacin administration on days 10 and 11 after hCG injection resulted in an increase in the LH/hCG **receptor** binding activity and rigidity of ovarian membrane lipids, as determined by **fluorescence polarization** of 1,6-diphenyl-1,3,5-hexatriene (DPH) probe. This effect was apparent within 7 days after indomethacin and ASA treatment. Both estradiol and. . . Unlike testosterone, the administration of dihydrotestosterone induced a decrease  
in membrane lipid rigidity and reduced the accessibility of the LH/hCG **receptor**. Inhibitors of prostaglandin F2alpha (PGF2alpha) synthesis, as the endogenous mediator of luteolysis, were shown to delay the regression of the. . . .

L2 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2002 ACS  
ACCESSION NUMBER: 1998:112498 CAPLUS  
DOCUMENT NUMBER: 128:176476  
TITLE: A method for quantitating competitive binding of molecules to **steroid** hormone **receptors** utilizing **fluorescence polarization**  
INVENTOR(S): Bolger, Randall E.; Ervin, Kerry M.; Lowery, Robert G.; Checovich, William J.  
PATENT ASSIGNEE(S): Panvera Corp., USA; Burke, Thomas, J.  
SOURCE: PCT Int. Appl., 37 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9805962	A1	19980212	WO 1997-US13538	19970801
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
PRIORITY APPLN. INFO.:			US 1996-23034P	P 19960802

TI A method for quantitating competitive binding of molecules to  
steroid hormone **receptors** utilizing **fluorescence**  
**polarization**

AB The system comprises mixing a **fluorescence**-emitting compd. that  
binds to the **steroid hormone receptors**, particularly  
the **estrogen receptor**, in a soln. contg. the **steroid**  
**hormone receptors**. Then, measuring the **fluorescence**  
**polarization** of the soln. Subsequently, incubating the soln. with  
at least one mol. that may compete with the compd. for interaction with  
the **steroid hormone receptors**. Measuring the  
**fluorescence polarization** of the soln. again. Finally,  
comparing the **fluorescence polarization** measurements  
to quantify any competitive interaction. A **fluorescence**  
-emitting compd. such as a **fluorescence**-emitting hormone can be  
used in combination with a fluorophore covalently coupled to an  
oligonucleotide to study how hormone and oligonucleotide binding to the  
hormone **receptor** are affected by each other.

ST **steroid receptor compd binding fluorescence**  
**polarization**

IT Nucleic acids  
RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)  
(**fluorescence**-labeled; method for quantitating competitive  
binding of mols., including nucleotides, to **steroid hormone**  
**receptors** utilizing **fluorescence polarization**  
)

IT DNA  
RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)  
(labeled with **fluorescein**; method for quantitating  
competitive binding of mols., including nucleotides, to **steroid**  
**hormone receptors** utilizing **fluorescence**  
**polarization**)

IT **Polarized fluorescence**  
(method for quantitating competitive binding of mols. to  
**steroid hormone receptors** utilizing  
**fluorescence polarization**)

IT **Estrogen receptors**  
Estrogens  
**Steroid receptors**  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(method for quantitating competitive binding of mols. to  
**steroid hormone receptors** utilizing  
**fluorescence polarization**)

IT DNA  
Nucleic acids  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(method for quantitating competitive binding of mols., including  
nucleotides, to **steroid hormone receptors** utilizing  
**fluorescence polarization**)

IT **Estrogen receptors**  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(recombinant; method for quantitating competitive binding of mols.,  
including nucleotides, to **steroid hormone receptors**  
utilizing **fluorescence polarization**)

IT 18930-97-7D, 5,6,11,12-Tetrahydrochrysene, derivs.  
RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)  
(**fluorescence** emitting hormone; method for quantitating  
competitive binding of mols. to **steroid hormone**  
**receptors** utilizing **fluorescence polarization**  
)

IT 50-28-2, Estradiol, biological studies  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL  
(Biological study); PROC (Process)  
(method for quantitating competitive binding of mols. to

steroid hormone receptors utilizing  
fluorescence polarization)  
IT 2321-07-5D, Fluorescein, DNA labeled with  
RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)  
(method for quantitating competitive binding of mols., including  
nucleotides, to steroid hormone receptors utilizing  
fluorescence polarization)

L2 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:395830 CAPLUS

DOCUMENT NUMBER: 127:107177

TITLE: Phospholipase C inhibitor, U73122, releases  
intracellular Ca<sup>2+</sup>, potentiates Ins(1,4,5)P<sub>3</sub>-mediated  
Ca<sup>2+</sup> release and directly activates ion channels in  
mouse pancreatic acinar cells

AUTHOR(S): Mogami, Hideo; Mills, Chris Lloyd; Gallacher, David  
V.

CORPORATE SOURCE: The Physiological Lab., Liverpool, L69 3BX, UK

SOURCE: Biochemical Journal (1997), 324(2), 645-651

CODEN: BIJOAK; ISSN: 0264-6021

PUBLISHER: Portland Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB It is recognized in many cellular systems that the **receptor**  
/G-protein activation of phospholipase C and Ins(1,4,5)P<sub>3</sub> prodn. is the  
transduction pathway regulating the release of Ca<sup>2+</sup> from internal stores.  
Ca<sup>2+</sup> signals can now be monitored at the level of single cells but the  
biochem. detection of Ins(1,4,5)P<sub>3</sub> cannot match this resoln. It is often  
difficult or impossible to directly attribute responses evoked in single  
cells by putative phospholipase C-coupled agonists to changes in  
Ins(1,4,5)P<sub>3</sub> levels. U 73122 is an amino **steroid** that is  
reported to act as a specific inhibitor of phospholipase C and it has  
become an important tool in establishing the link between phospholipase C  
activation and cellular Ca<sup>2+</sup> signaling. In the present study we use both  
patch-clamp electrophysiol. and the imaging of **fluorescent** Ca<sup>2+</sup>  
indicators to investigate the effect of U 73122 in mouse pancreatic

acinar

cells. The study reveals that U 73122 has effects other than the  
inhibition of phospholipase C. U 73122 can directly activate ion  
channels. It can itself promote the release of Ca<sup>2+</sup> from intracellular  
stores in permeabilized cells and in intact cells it triggers a release

of

Ca<sup>2+</sup> that is initiated specifically at the secretory pole of these  
morphol. and functionally **polarized** cells. We also present  
evidence that U 73122 can potentiate the response to Ins(1,4,5)P<sub>3</sub>; this

is

seen both in permeabilized cells and in patch-clamp protocols in which  
cells are internally dialyzed with submaximal concns. of Ins(1,4,5)P<sub>3</sub>.  
The effects of U 73122 are therefore multiple and not specific for the  
inhibition of phospholipase C. Importantly, all the effects described  
influence Ca<sup>2+</sup> signaling yet in many exptl. protocols some of these  
effects can go unnoticed and might in error be attributed simply to the  
inhibition of Ins(1,4,5)P<sub>3</sub> prodn.

L2 ANSWER 5 OF 8

MEDLINE

DUPLICATE 2

ACCESSION NUMBER: 86221226 MEDLINE

DOCUMENT NUMBER: 86221226 PubMed ID: 3011559

TITLE: Sex steroid and prostaglandin interactions upon the  
purified rat myometrial plasma membranes.

AUTHOR: Delicostantinos G; Fotiou S

SOURCE: MOLECULAR AND CELLULAR ENDOCRINOLOGY, (1986 May) 45 (2-3)  
149-56.

Journal code: 7500844. ISSN: 0303-7207.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals  
ENTRY MONTH: 198607  
ENTRY DATE: Entered STN: 19900321  
Last Updated on STN: 19900321  
Entered Medline: 19860710

AB . . . concentration of  $1 \times 10^{-6}$  M for 1 h at 37 degrees C, bind into MPM at pmolar concentrations. Unlabeled **steroids** inhibited [3H]PGE2 and [3H]PGF2 alpha binding to MPM in a dose-dependent manner. Membrane-bound and free **steroids** or PGs were found to be essentially unchanged under the present incubation conditions.  $\text{Ca}^{2+}$  ions up to 10 mM increased **steroid** binding into MPM. Molecular interactions between **steroids** and MPM were assessed by measuring the steady-state **fluorescence polarization** of 1,6-diphenyl-1,3,5-hexatriene (DPH), and by estimating the changes in the allosteric properties of MPM-bound ( $\text{Na}^+ + \text{K}^+$ )ATPase by fluoride ( $\text{F}^-$ ). **Steroids** appear to increase the MPM fluidity, evaluated through changes in the Hill coefficient for MPM-bound ( $\text{Na}^+ + \text{K}^+$ )ATPase by  $\text{F}^-$  and by the **fluorescence polarization** method. Binding of sex **steroids** to MPM increased the membrane fluidity and decreased the binding of the uterus stimulatory PGs by membrane **receptors**. These studies provide a basis for postulating that a 'non-genomic' mechanism of sex **steroids** induces reduction of uterine contractions.

L2 ANSWER 6 OF 8 MEDLINE

ACCESSION NUMBER: 77159867 MEDLINE  
DOCUMENT NUMBER: 77159867 PubMed ID: 856460  
TITLE:

Fluidity of membrane lipids and lateral mobility of concanavalin A receptors in the cell surface of normal lymphocytes and lymphocytes from patients with malignant lymphomas and leukemias.

AUTHOR: Ben-Bassat H; Polliak A; Rosenbaum S M; Naparstek E; Shouval D; Inbar M

SOURCE: CANCER RESEARCH, (1977 May) 37 (5) 1307-12.  
Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 197706

ENTRY DATE: Entered STN: 19900313  
Last Updated on STN: 19900313  
Entered Medline: 19770622

AB . . . with nonmalignant and malignant disorders were studied for fluidity of membrane lipids and lateral mobility of concanavalin A (Con

A)

**receptors**. The degree of fluidity of the surface membrane lipid core was monitored quantitatively by **fluorescence polarization** analysis using the probe 1,6-diphenyl-1,3,5-hexatriene embedded in lipid regions of the surface membrane of intact cells. Mobility of Con A surface **receptors** was determined by the cap-forming ability after binding of **fluorescent** Con A. The present studies were performed on lymphocytes from 28 patients with malignant lymphomas, 22 patients with leukemia, 28. . . fluidity was less pronounced in lymphocytes isolated from leukemic patients in

clinical

remission and from leukemic patients receiving treatment with **steroids**. The results also show a marked difference in the cap-forming ability of lymphocytes from patients with malignant lymphomas or leukemia. . . a higher cap-forming ability. The cap-forming ability of cells from patients with chronic lymphocytic leukemia was unaffected

by

treatment with **steroids**. The present results, which are in line with previous observations, have shown that normal lymphocytes can be characterized by a low degree of lipid fluidity but a high degree of mobility of Con A **receptors**, whereas leukemic lymphocytes are

characterized by a high degree of lipid fluidity but a low degree of mobility of **Con A receptors**. These results confirmed our general hypothesis on the dynamic interrelation between membrane lipids and membrane protein **receptors**, and they indicate that the widely accepted term "membrane fluidity" requires better consideration for different membrane components.

L2 ANSWER 7 OF 8 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.  
ACCESSION NUMBER: 78105430 EMBASE  
DOCUMENT NUMBER: 1978105430  
TITLE: Fluidity of membrane lipids and lateral mobility of concanavalin A receptors in the cell surface of normal lymphocytes and lymphocytes from patients with malignant lymphomas and leukemias.  
AUTHOR: Ben Bassat H.; Polliak A.; Rosenbaum S.M.; et al.  
CORPORATE SOURCE: Dept. Hematol. Med. A, Chanock Cent. Virol., Hebrew Univ. Hadassah Med. Sch., Jerusalem, Israel  
SOURCE: Cancer Research, (1977) 37/5 (1307-1312).  
CODEN: CNREA8  
COUNTRY: United States  
DOCUMENT TYPE: Journal  
FILE SEGMENT: 016 Cancer  
025 Hematology  
005 General Pathology and Pathological Anatomy  
026 Immunology, Serology and Transplantation

LANGUAGE: English

AB . . . with nonmalignant and malignant disorders were studied for fluidity of membrane lipids and lateral mobility of concanavalin A (Con A)

**receptors**. The degree of fluidity of the surface membrane lipid core was monitored quantitatively by **fluorescence polarization** analysis using the probe 1,6-diphenyl-1,3,5-hexatriene embedded in lipid regions of the surface membrane of intact cells. Mobility of Con A surface **receptors** was determined by the cap-forming ability after binding of **fluorescent** Con A. The present studies were performed on lymphocytes from 28 patients with malignant lymphomas, 22 patients with leukemia, 28. . . fluidity was less pronounced in lymphocytes isolated from leukemic patients in

clinical

remission and from leukemic patients receiving treatment with **steroids**. The results also show a marked difference in the cap-forming ability of lymphocytes from patients with malignant lymphomas or leukemia. . . a higher cap-forming ability. The cap-forming ability of cells from patients with chronic lymphocytic leukemia was unaffected

by

treatment with **steroids**. The present results, which are in line with previous observations, have shown that normal lymphocytes can be characterized by a low degree of lipid fluidity but a high degree of mobility of Con A **receptors**, whereas leukemic lymphocytes are characterized by a high degree of lipid fluidity but a low degree of mobility of Con A **receptors**. These results confirmed the general hypothesis on the dynamic interrelation between membrane lipids and membrane protein **receptors**, and they indicate that the widely accepted term 'membrane fluidity' required better consideration for different membrane components.

L2 ANSWER 8 OF 8 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.  
ACCESSION NUMBER: 1977:205439 BIOSIS  
DOCUMENT NUMBER: BA64:27803  
TITLE: FLUIDITY OF MEMBRANE LIPIDS AND LATERAL MOBILITY OF CONCAVALIN A RECEPTORS IN THE CELL SURFACE OF NORMAL LYMPHOCYTES AND LYMPHOCYTES FROM PATIENTS WITH MALIGNANT LYMPHOMAS AND LEUKEMIAS.  
AUTHOR(S): BEN-BASSAT H; POLLIACK A; ROSENBAUM S M; NAPARSTEK E; SHOUVAL D; INBAR M  
SOURCE: CANCER RES, (1977) 37 (5), 1207-1312.



FILE SEGMENT:

BA; OLD

LANGUAGE:

Unavailable

AB. . . with nonmalignant and malignant disorders were studied for fluidity of membrane lipids and lateral mobility of concanavalin A (Con A) **receptors**. The degree of fluidity of the surface membrane lipid core was monitored quantitatively by **fluorescence polarization** analysis using the probe 1,6-diphenyl-1,3,5-hexatriene embedded in lipid regions of the surface membrane of intact cells. Mobility of Con A surface **receptors** was determined by the cap-forming ability after binding of **fluorescent** Con A. The present studies were performed on lymphocytes from 28 patients with malignant lymphomas, 22 patients with leukemia, 28. . . membrane fluidity was less pronounced in lymphocytes isolated from leukemic patients in clinical remission and leukemic patients receiving treatment with **steroids**. The results also show a marked difference in the cap-forming ability of lymphocytes from patients with malignant lymphomas or leukemia. . . a higher cap-forming ability. The cap-forming ability of cells from patients with chronic lymphocytic leukemia was unaffected

by

treatment with **steroids**. The present results, which are in line with previous observations, showed that normal lymphocytes can be characterized by a low degree of lipid fluidity but a high degree of mobility of Con A **receptors**, whereas leukemic lymphocytes are characterized by a high degree of lipid fluidity but a low degree of mobility of Con A **receptors**. These results confirmed the general hypothesis on the dynamic interrelation between membrane lipids and membrane protein **receptors**, and they indicate that the widely accepted term membrane fluidity requires better consideration for different membrane components.

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